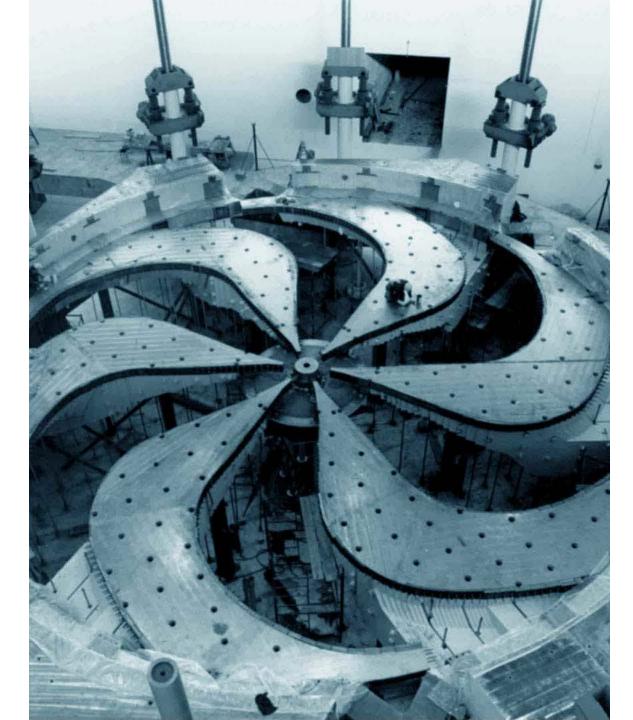


Production of medical radionuclides at TRIUMF

Valery Radchenko, PhD TRIUMF/UBC

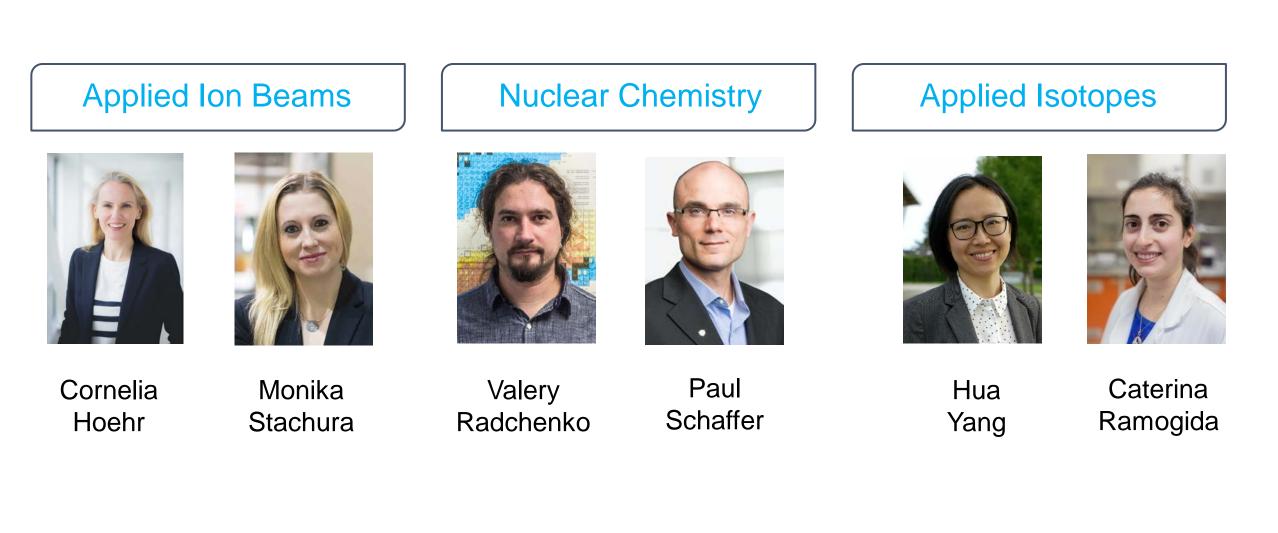
July 31st, 2023



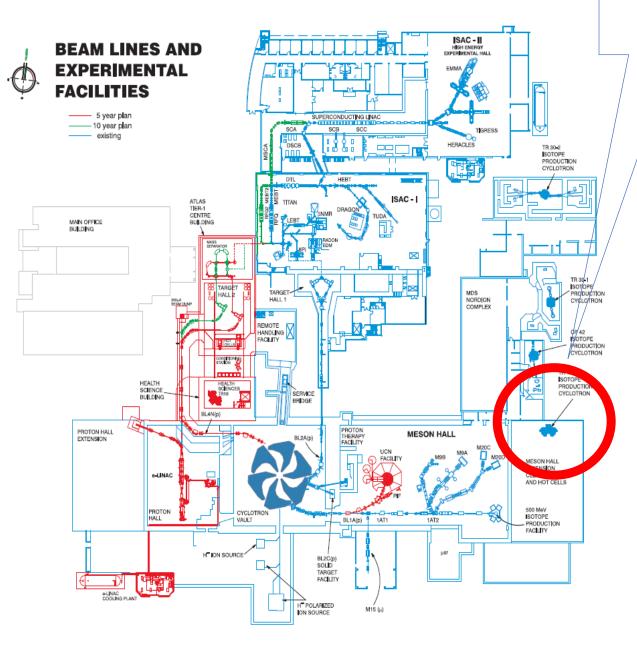
Discovery, accelerate

1

2023-07-30



Production of medical radionuclides at TRIUMF:TR-13



ISAC

TR-13

Production of pre-clinical quantitates of imaging and therapeutic radionuclides

Isotope Production Facility (IPF), BL1A

ARIEL

Collection chamber Proton and electron beamlines

IAMI TR-24

TR-13: work horse of Life Sciences





TR-13 Operation team:

David Prevost

Spencer Staiger

Ryley Morgan

Toni Epp

Routine production of ¹⁸**F** and ¹¹**C** for clinical collaboration partners (UBC hospital, etc.)

Production of diagnostic radionuclides:

Liquid targets: ⁶⁸Ga, ⁴⁴Sc, ⁸⁶Y, ⁸⁹Zr..... Easy to handle/transport Usually, lower yield compared to solid targets

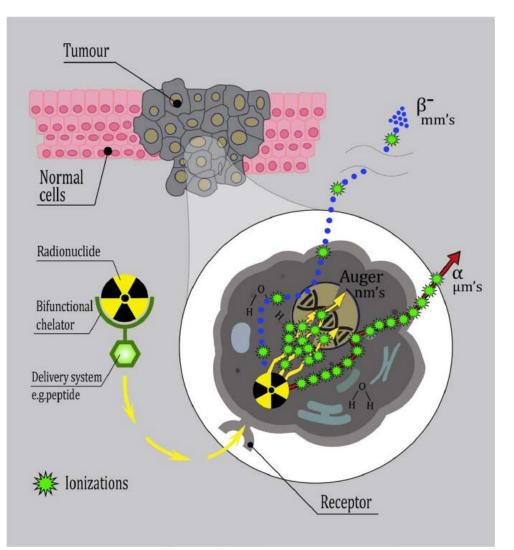
Solid targets: ⁶⁸Ga, ⁴⁴Sc, ⁴⁵Ti, ⁶⁴Cu, ⁸⁶Y, ⁹⁰Nb, ⁸⁹Zr, ¹⁵⁵Tb,.....

Production of therapeutic radionuclides:

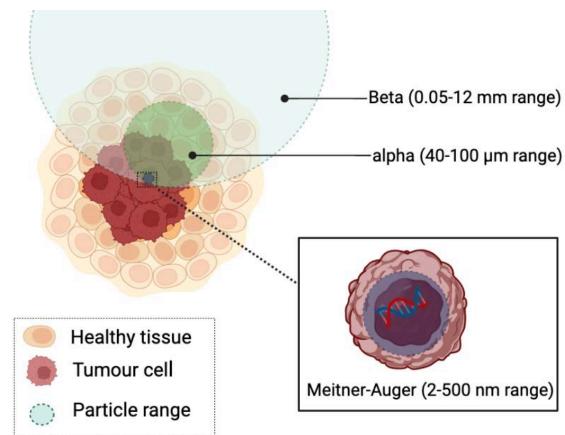
^{197m+g}Hg, ¹¹⁹Sb, ¹⁰³Pd, ¹³⁵La and ¹⁶⁵Er

Targeted Radiation Therapy (TRT): Meitner-Auger Emitters

Radiopharmaceutical design

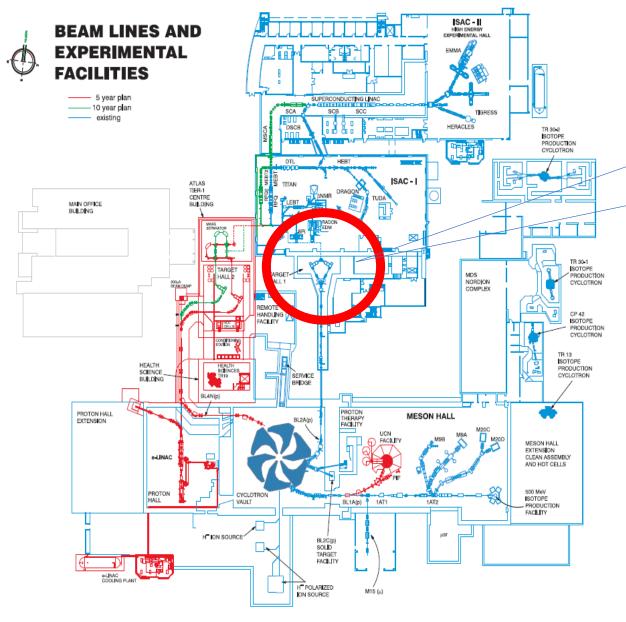


Beta vs. Alpha vs. Auger Emitters



V. Radchenko and C. Hoehr, Nucl. Phys. News, 2020 Filosofov D, Kurakina E, Radchenko V. Nucl. Med. Biol. 2021

Production of medical radionuclides at TRIUMF:ISAC



TR-13

ISAC

Production of pre-clinical quantitates of imaging and therapeutic radionuclides

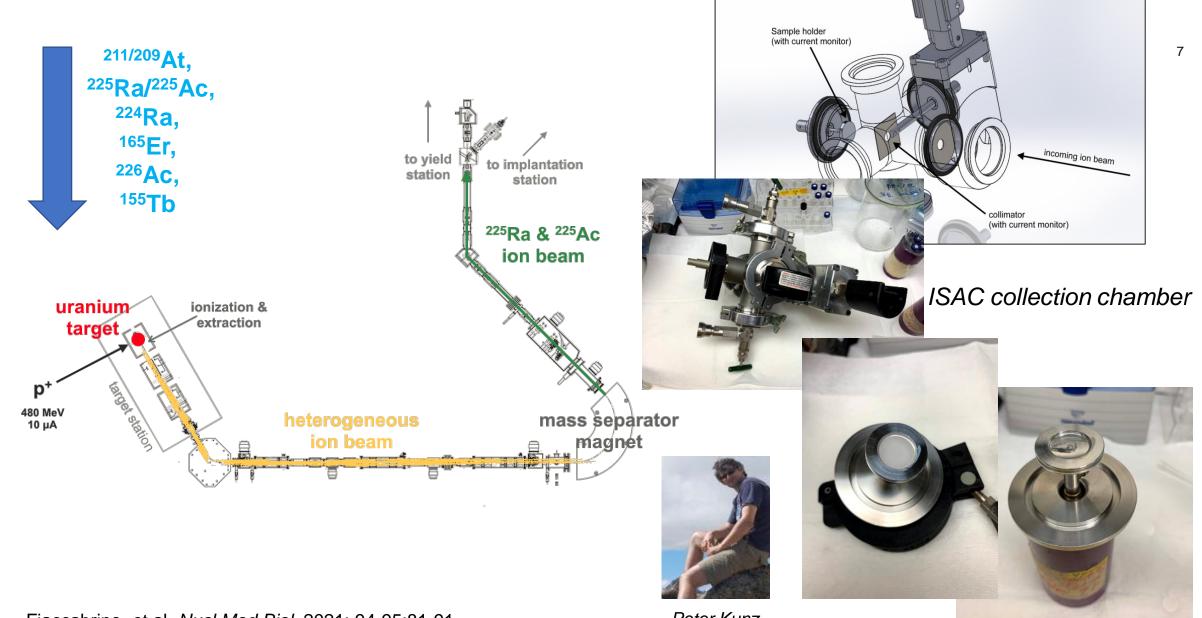
Isotope Production Facility (IPF), BL1A

ARIEL

Collection chamber Proton and electron beamlines

Isotope Separation On-Line

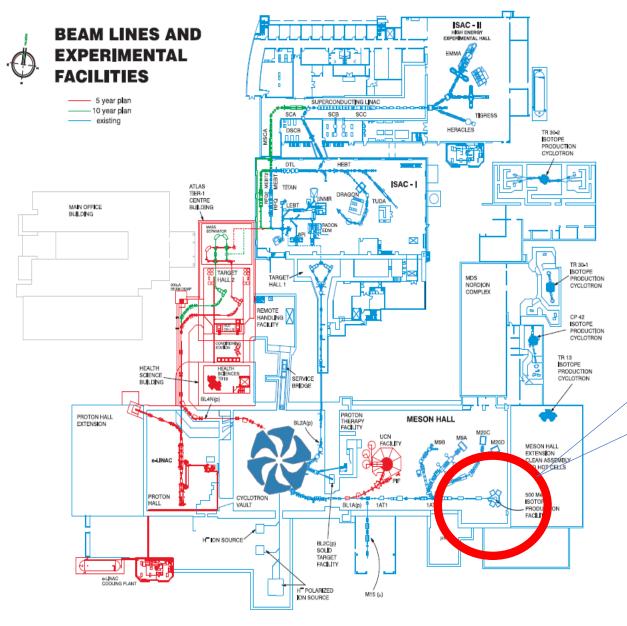
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Fiaccabrino, et al. Nucl Med Biol. 2021; 94-95:81-91.

Peter Kunz

Production of medical radionuclides at TRIUMF:BL1A



TR-13

ISAC

Isotope Production Facility (IPF), BL1A

Large scale production of therapeutic radionuclides

Collection chamber Proton and electron beamlines

E

²²⁵Ac/²¹³Bi promising system for Targeted Alpha Therapy (TAT)

225Ra B: 14.9 d 225Ac a: 10.0 d 221Fr a: 4.9 m 217At a: 32.3 ms 213Bi β: 45.6 m a: 45.6 m 98% 209**TI** 213Po α: 4.2 µs 6: 2.2 m 209Pb β: 3.3 h 209**Bi**

http://www.nndc.bnl.gov/chart/

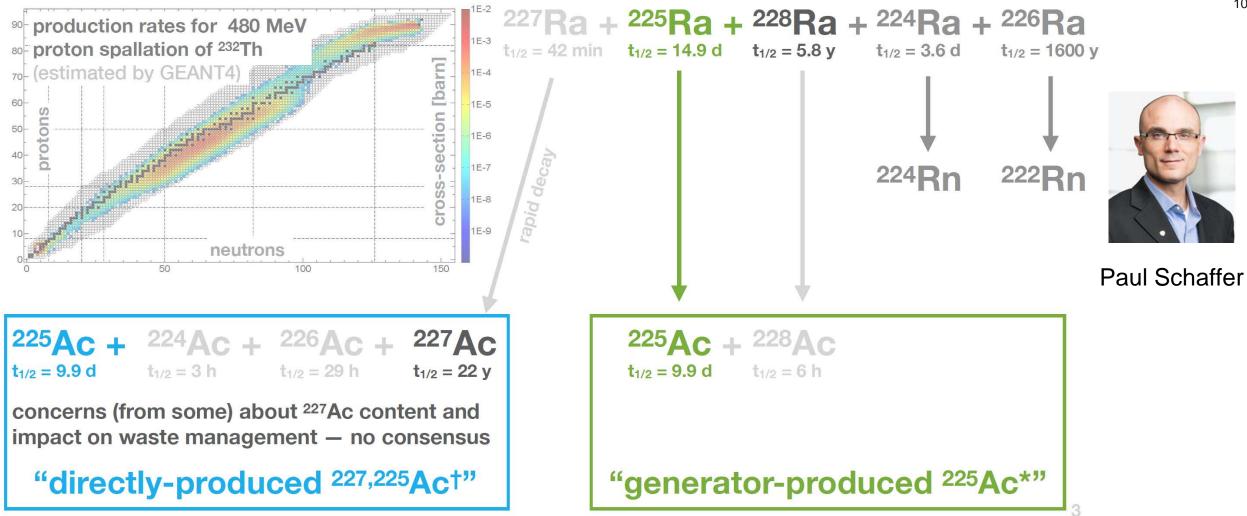
²²⁵Ac (t_{1/2} 9.92 d) in combination with specific biomolecules (e.g. peptides) is a promising system for Targeted Alpha Therapy (TAT)

- ²²⁵Ac/²¹³Bi (t_{1/2} 45.59 min) generator system provides
 accolorator independent source of ²¹³Bi for modical appli
 - accelerator independent source of ²¹³Bi for medical applications

• Supply is limited/challenging for clinical translation

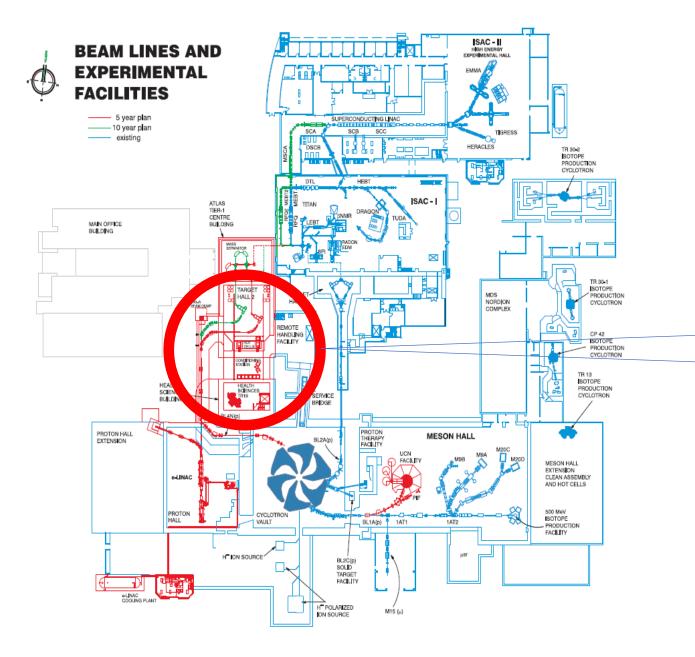
International Atomic Energy Agency. Technical Meeting Report "Alpha Emitting Radionuclides and Radiopharmaceuticals for Therapy" IAEA Headquarters Vienna, Austria. 24-28 June **2013**

High-purity ²²⁵Ac production available via ²²⁵Ra



10

Production of medical radionuclides at TRIUMF: ARIEL



TR-13

ISAC

Isotope Production Facility (IPF), BL1A

ARIEL

Proton and electron beamlines

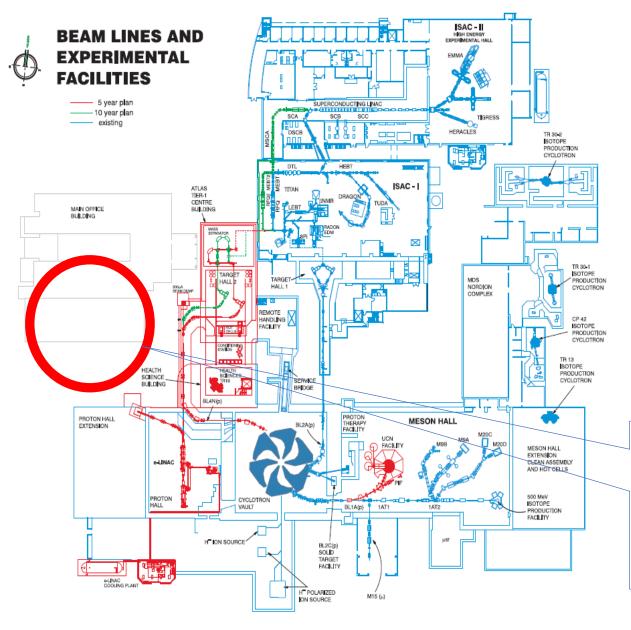
IAMI TR-24

Proton-induced ²²⁵Ac production at ARIEL: proton beamline

Proton Energy Distribution • Hundreds of co-produced isotopes including, ²²⁵Ac, ²²⁴Ra, ²²³Ra, ²¹³Bi, ²¹²Pb, ²¹²Bi 800 Protons in primary proton beam Protons in ISOL target *1.4 Protons in proton beam dump *10 700 Production in 6 cm diameter, 5 cm thick, 1.5 kg Th metal, [10 µA⁻¹ · s⁻¹] Proton Fluence [arb. units] in beam 90 · in ISOL target 1013 40 600 ISOL target beam dump 80 10-1 012 70 Th target 500 units in beam dump 10-1011 60 Suctors 50 40 (70% of beam) [arb 400 10¹⁰ Ę 40 nten 10⁹ tra -20 300 30 10-5 primary 20 -40 proton beam 200 120 140 160 100 100 20 40 60 80 100 120 140 0 longitudinal position [cm] Neutrons Irradiation Delivery to ARIEL Pneumatic distribution 0 100 200 300 400 500 0 position hot cell system Proton Energy [MeV] Place symbiotic medical target in-between 40 CEM ²³²Th(p,X)²²⁵Ac ISOL target and beam dump Rertin 35 Cross Section [mb] • Pneumatic target delivery system from 30 25 ARIEL hot cell complex to proton target 20 station Zhuikov, et al. 2011 15 Ermolaev, et al. 2012 • Independent operation from ISOL target This work 10 development for in-cell processes and 5 irradiation station required 0 25 50 100 125 150 175 200 0 75 Proton Energy [MeV]

J.W. Weidner et al., Applied Radiation and Isotopes 70 (2012) 2602-2607

Production of medical radionuclides at TRIUMF



TR-13

ISAC

Isotope Production Facility (IPF), BL1A

ARIEL

Collection chamber Proton and electron beamlines

IAMI TR-24 GMP facility

Institute for Advanced Medical Isotopes (IAMI)

IAMI will unite interdisciplinary partners to create a world-class center for advanced isotope research, development, and production for the life sciences.

TR-24: Proton Energy 24 MeV, 500 μ A will enable (p,2n) along with (p, n) reactions





IAMI Founding Partners



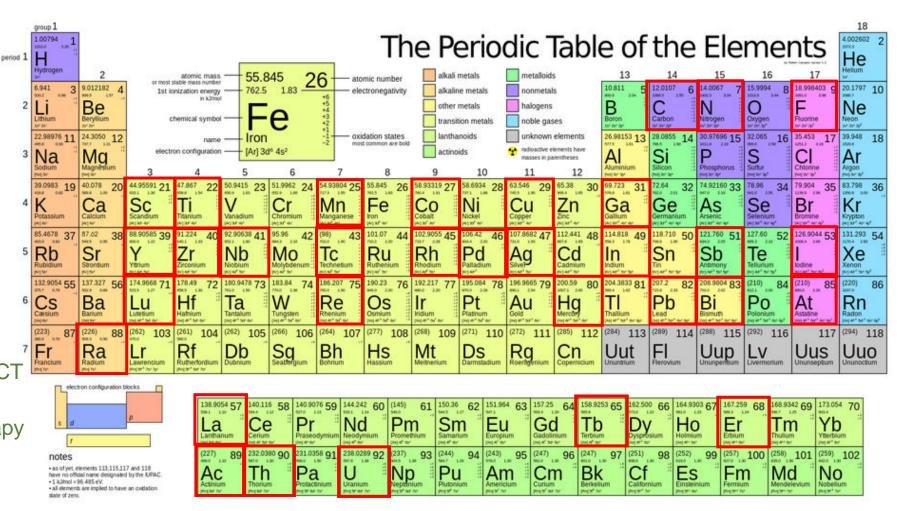




¹¹C PET

- ¹⁸F PET
- ⁴⁴Sc PET/ ⁴⁷Sc β ⁻ therapy
- ⁴⁵Ti PET
- ^{52,54}Mn PET
- ⁵⁵Co PET
- ⁶⁴Cu PET/ ⁶⁷Cu β⁻ therapy
- ⁶⁸Ga PET/ ⁶⁷Ga Auger therapy
- ⁸⁶Y PET/⁹⁰Y β⁻ therapy
- ⁸⁹Zr PET
- ⁹⁰Nb PET
- 99mTc SPECT/ 94mTc PET
- ^{103m}Rh Auger therapy
- ¹¹¹In SPECT
- ¹¹⁹Sb Auger/¹¹⁸Sb PET/¹¹⁷Sb SPECT
 ¹²⁴I PET/ ¹²⁵I Auger therapy
 ¹⁴⁹Tb Alpha therapy/ ¹⁶¹Tb β⁻ therapy
- ¹⁶⁵Er Auger therapy
- ¹⁷⁷Lu β^{-} therapy
- ²⁰³Pb/ ²¹²Pb Alpha therapy
- ²¹³Bi Alpha therapy
- ^{223,224}Ra Alpha therapy
- ²²⁵Ac Alpha therapy^{227,228}Th Alpha therapy

TRIUMF's "Hot Kitchen"



Blue available Green can/planning be produced Yellow commercially available ISAC and Th spallation provides endless possibility for production of many other medical isotopes

Summary

TR-13

- Supporting clinical collaborations with radiohalogens (¹⁸F and ¹¹C)
- Production of emerging radiometals for PET
- Production of promising therapeutic radionuclides for Auger therapy ISAC
- Production of pre-clinical quantities of many novel/promising imaging and therapeutic radionuclides

IPF (BL1A)

- Large scale (clinical relevant) production of emerging therapeutic radionuclides (²²⁵Ac)
 ARIEL
- Proton beamline for pre-clinical and clinical supply of therapeutic radionuclides
- Possibility for electron beamline, utilization of (γ, n) (γ, p) reactions for production of medical radionuclides

IAMI

- Clinical level supply of medical radionuclides (¹⁸F, ^{99m}Tc, ⁸⁹Zr, ⁶⁴Cu....) and GMP formulation of radiopharmaceuticals
- Provide more flexibility (e.g. enable p, 2n)

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%TRIUMF

Gokce Engudar

Siuwong Chan

Viktoria Krol

TR-13 team

Toni Epp

Syddansk Universitet

Dave Prevost

Ryley Morgan

Helge Thisgaard

Lorraine Ge Gaenaelle

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Aivija Grundmane

Parmissa Rhandhawa

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Raymond Reilly Zhongli Cai Noor Alsaden Constantine Georgiou

British Columbia Cancer Agency

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UBC chemistry Orvig lab

JINR

Dmitry Filosofov LNP Radiochemistry group Elena Kurakina

UW Madison

Jonathan W. Engle Todd Barnhart Aeli Olson Cyclotron group





Canadian Société Cancer canadienne Society du cancer



Thank you Merci



vradchenko@trumf.ca

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